Figure 1 - p-RNA Structure

Pyranosyl-RNA

Advantageous properties:

Watson-Crick pairing mode exclusively

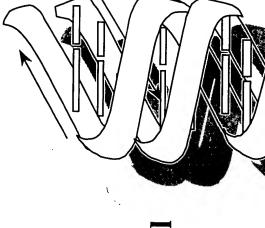
Antiparallel strand orientation exclusively

- Duplexes *more stable* than in DNA or RNA
 - Duplexes have a quasi-linear ladder structure
 - Pairing is enantioselective
 - Potential to replicate *without enzymes* (thus prebiotic ancestor to RNA?)
 - No pairing with DNA or RNA

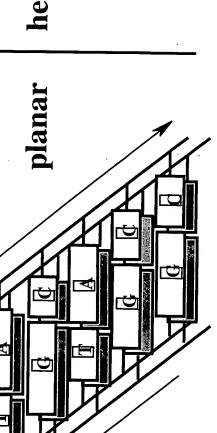
Figure 2 - P-RNA Planar and DNA Helical designations with the second s

p-RNA

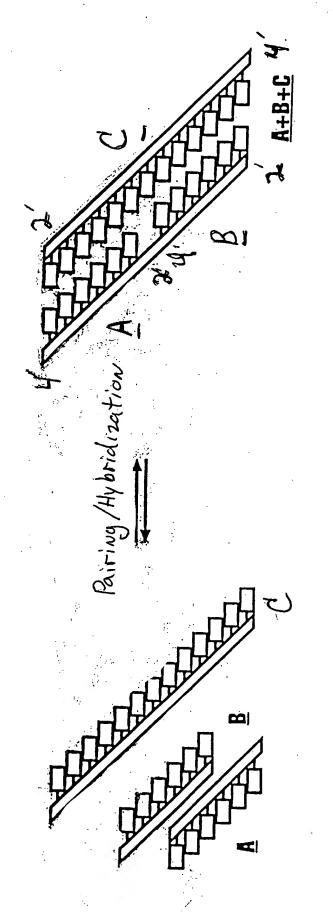
DNA / RNA



helical



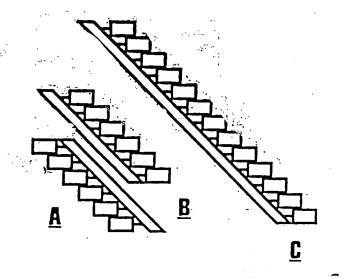
P-RNA Components A, B, and C l'igure 3-



P-RNA Components A, B, and C UN-paired

P-RNA Components A, B, and C Paired or hybridized

Figure 4- p-RNA Peptide Derivatized A, B, + C Components

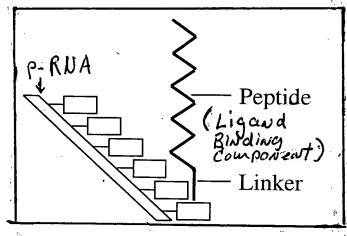


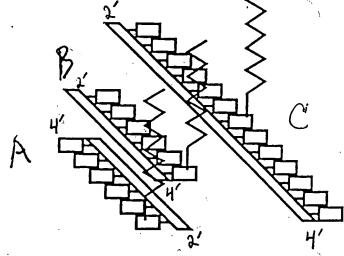
P-RNA Components

A, B, and C

(Programable Pairing Components)

Linking of
peptide library
with p-RNA
Components A, B, + C





P-RNA Peptide

derivatized

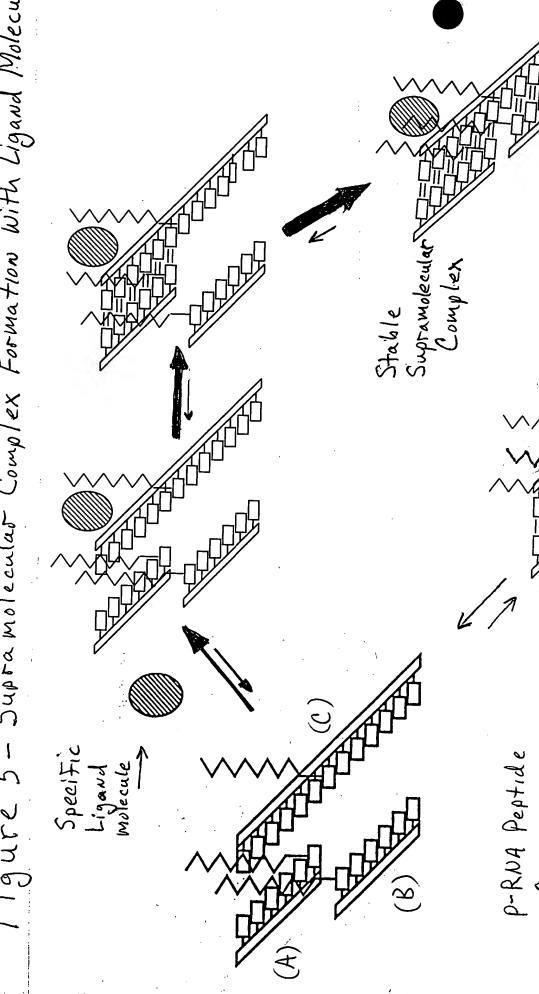
A, B, and C

Components

(Intermolecular Ligard Binding Components)

dented the state of the state o

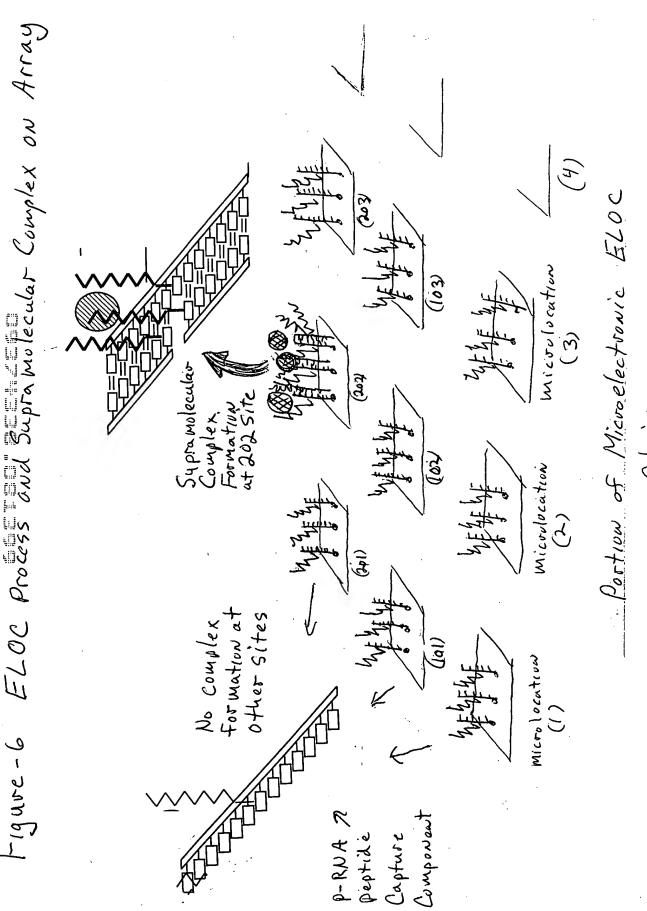
Figure 5- Supramolecular Complex Formation With Ligand Molecule der ger ger generalische gein gelig bei erzig gemein gener B. ge gemeiner gener Bande Rauft fraus der Grant Band Grant Grant



P-RNA Peptide Components A, B, + C

Stacker Structure with (No ligand molecule)

ELOC Process and F19 ure - 6



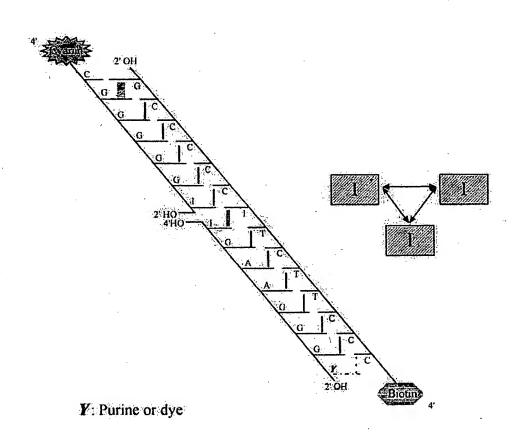
Ch. P

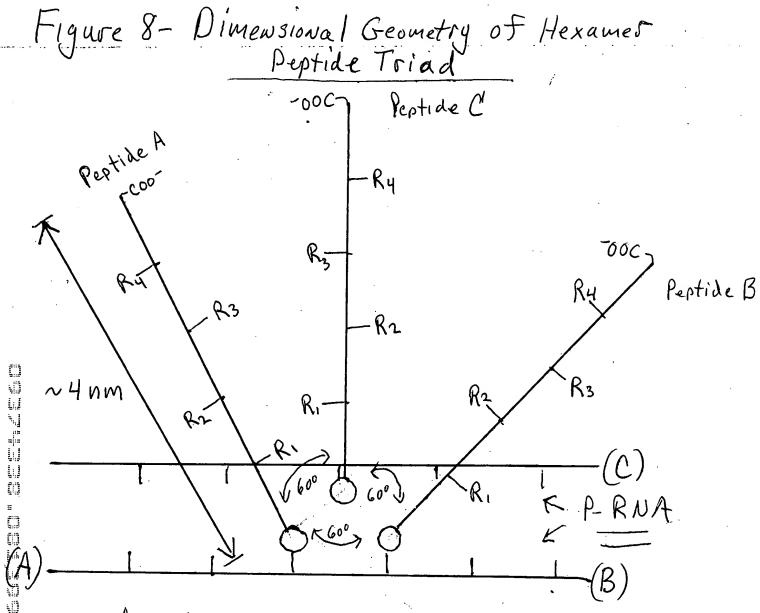
Figure 7 - Fully hybridized 7-mes p-RNA A, 7-mes p-RNAB, and complementary capture P-RNA C.

Oligo 90: 4' IGAAGGGY 2'

Oligo Cy91: 4' Cyanine-CGGGGGI 2'

Oligo Biot92: 4' Biotine-CCCTTCTICCCCCG 2'





Amino Acid R Groups
R1 ~ 0.6 nm
R2 ~ 0.96 nm
R3 ~ 1.3 nm
R4 ~ 1.7 nm

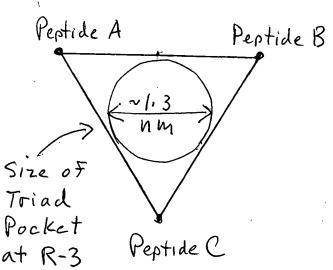
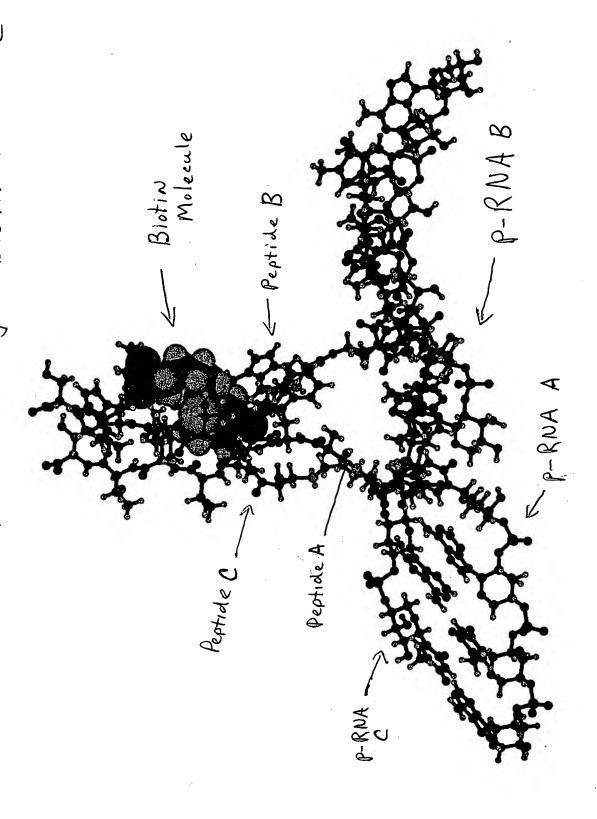
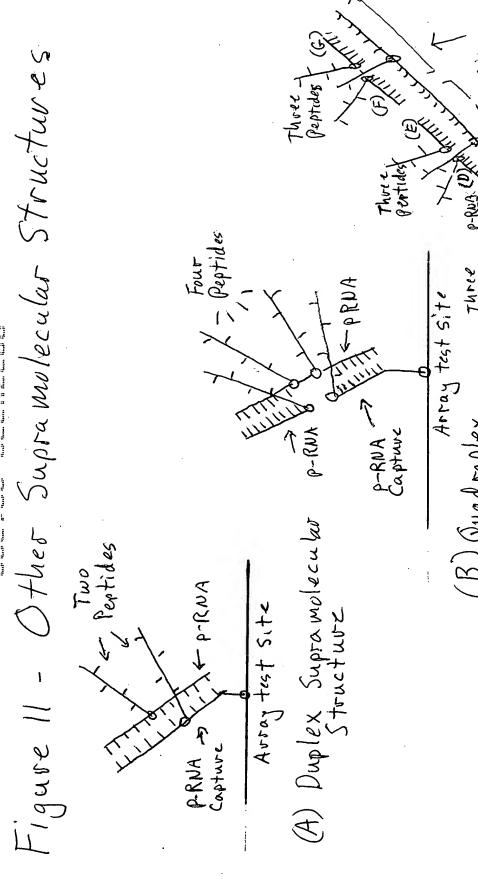


Figure 9 - p-RNA-Peptide Triad Suprastructure Complex Binding a Bistin Molecule



Binding Site for Acetylaholine (C) Un Favorable Peptide Triad Peptides with net position charges Tiqure 10 - Favorable and Un Favorable Peptide Triads P-RNA No binding For Binding Arety/Choline MICROGERAY (A) Acetylcholine Molecule The state of the s Acetyleholine CF. O. C. CF. Molecule Binding Site For Acetylcholine Boun d (B) Favorable Peptide Triad Micoro array Test SiteD Peptides . with met Negative charges

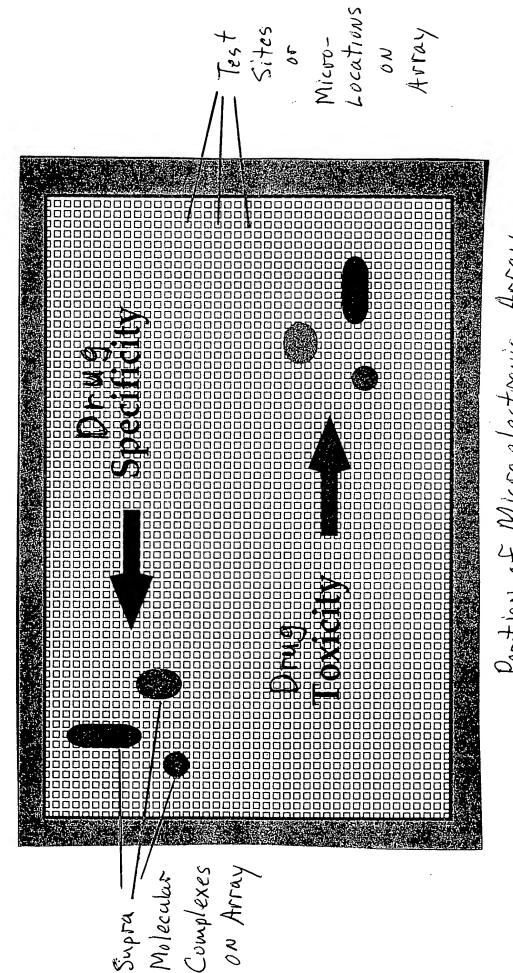


P-RNA Supra Molecular (B) Quadraplex Structure

(C) Multiplex Tirlad Supramolecular Horay test Site Structures

- P-RNA Capture

Jescriptor Array dan a familian dan dan dan Molecular 3 **[**..] į.) . . 1,9 ure 12



Portion of Micro electioni

Electronic Pertur Bation Improve Ligard Binding + 19 ure 13

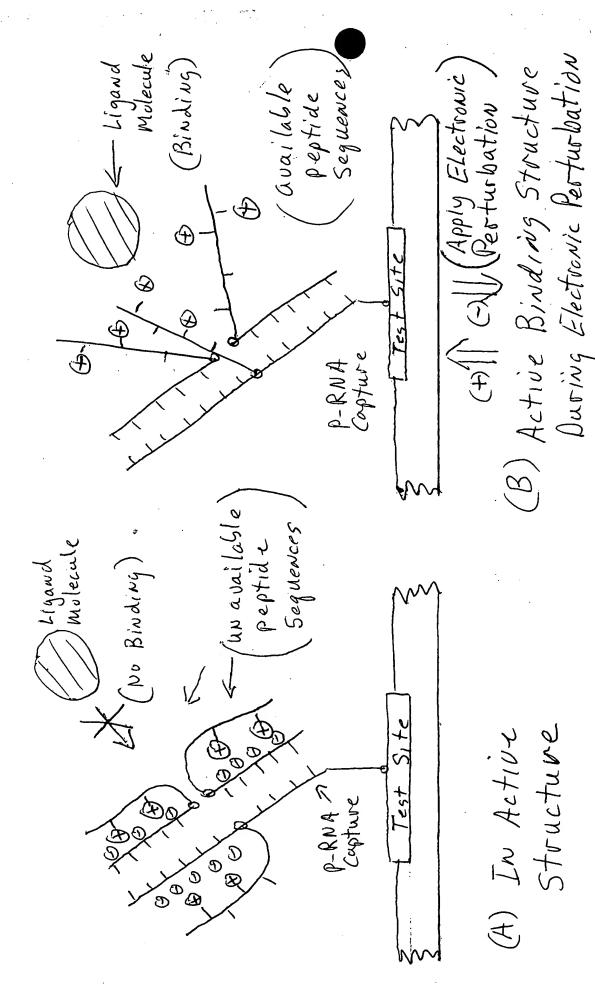


Figure 14- Specific p-RNA Hybridiza from on 10,000 Site Microelectronic Array

である。ことである。ことである。ことできる。 では、ことでは、他のできる。ことできる。 では、ことでは、他のできる。ことできる。 では、ことできる。ことできる。ことできる。 をは、ことできる。ことできる。ことできる。 をは、ことできる。

30 MICO

test Sites

Capture (92) Hybridized with

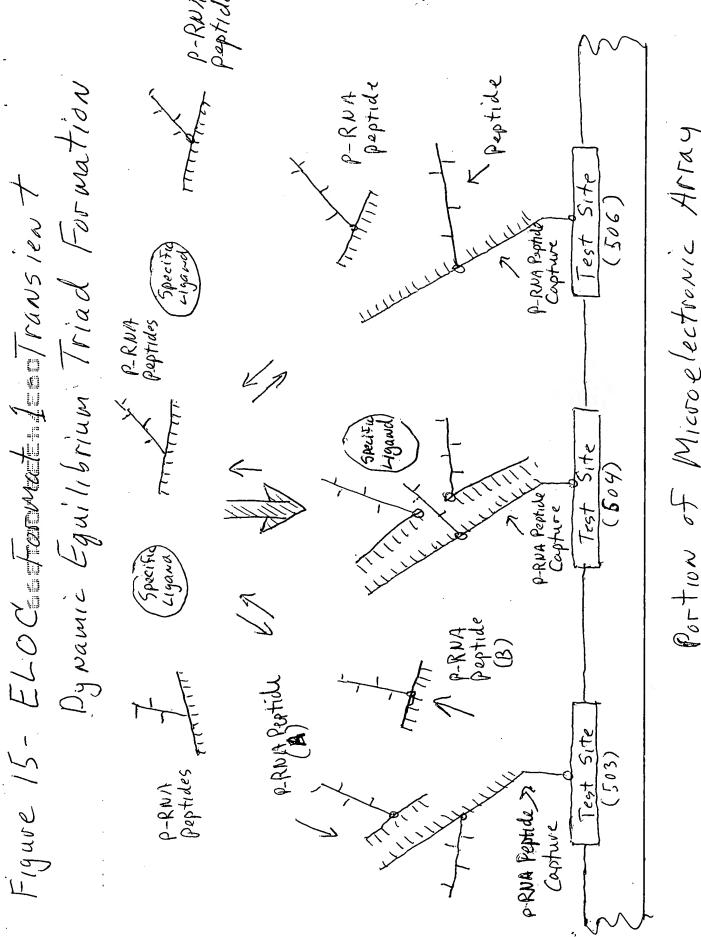
Specific P-RNA

P-RNA 91-CY3

Now-Specific Capture P-RNA 72

Portion of 10,000 site Microelectronic

Array



Portion of Microelectronic

ELOS Formation Process in Sclution Initial Complex Formation 1-19uve 16 Specific Linguid

Micro electionic Array

Final Supra molecular Complex Formation on Array

> Micro electronic Hrray

1

Flywire 17 - ELOG Frank Matton Heterogeneous Triad Formation Process

(A) Initial Formation of all supramolecular Structures

PRINT PORTURE STATE STAT

Add Specific Ligands

Desitic Ligand Ligand Lumplex Formation

Complex Formation with Specific

Microelectronic Array

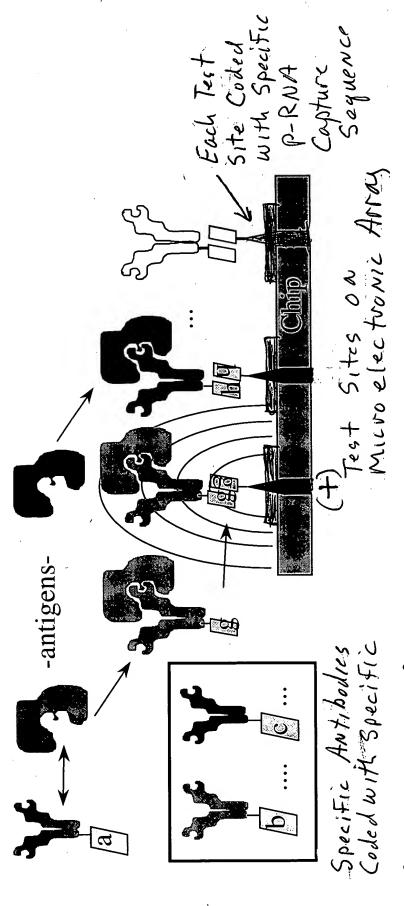
(B) Secondary
Formation of
Supra molecular
Complexes on Array

Microelectronic Auray

100

Multiplex Pertide Triads A's Synthetic Antibodies For Immuno Assay Secondary Flyorecen 1 Microelectronic Arrow X SOLL Test Site Specific Complementary P-RNA Capture to Specific P-RNA Complementury Capture or P-RNA Sequence Array Figure Specific Bound

#3 #4 Figure-19 - Modular Lumana Assay Using P-RNA Antibody Conjugates on Microelectronic Array



Complementary P-RNA Sequences to those Coded on the Array Molecular Weight = 3241.61

Exact Mass = 3239.73

Molecular Formula = C119H146N36O55P7S2

Mw calc: 3241.61

Mw found: 3243.3 ± 0.5 (error~ 0.4%)

Molecular Weight = 3241.61

Exact Mass = 3239.73

Molecular Formula = C119H146N36O55P7S2

Mw calc: 3241.61

Mw found: 3243.3 ± 0.5 (error~ 0.4%)

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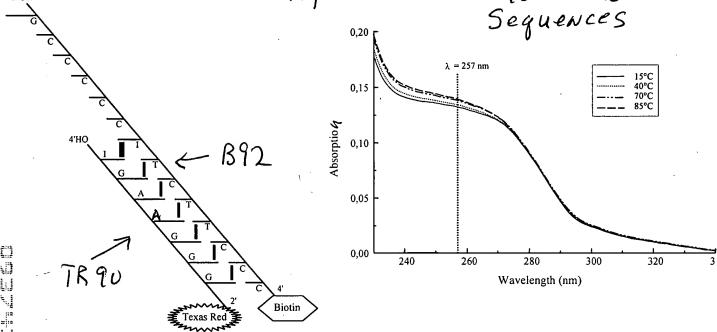
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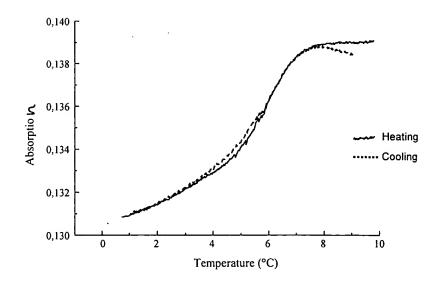
990566es1#13-15 RT: 0,49-0,56 AV: 3 NL: 6,96E3 F: - p ms [500,00-2000,00] 1620,7 100-90-80-70-Relative Abundance 60-50-40-1080.1 30-20-1631,3 10-1128,1 1009,7 785.9 651 0 0 800 600 1000 1200 1400 1600

Oligo TR90: 4' IGAAGGG-TexasRed 2' Oligo B92: 4' Biotin-CCCTTCTICCCCCG 2'

Figure 21- Structure for Hybridized TR-90 and B92 Sequences

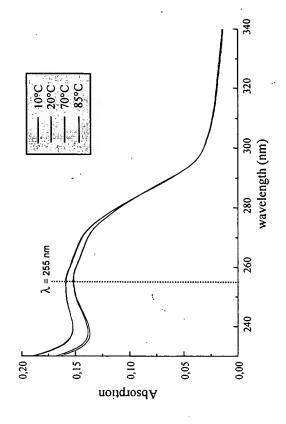


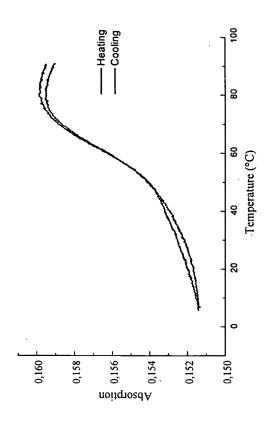
 $T_m = 59 \,^{\circ}\text{C}$ $Tris/HCl \, 0.01M \, ; \, pH \, 7$ $0.15 \, M \, NaCl$ $c = 5 \, \mu M + 5 \, \mu M$

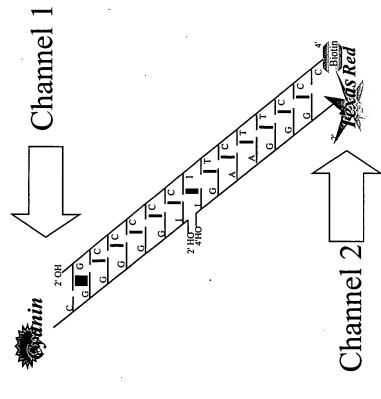


Oligo TR-90: 4" Cyanine3-CGGGGGI 2" Oligo Biot-92: 4" Biotin-CCCTTCTICCCCCG 2"

Figure 22.
Stracture for Hybridized TR90,
C43-91, and B92
P-RNA Conjugates

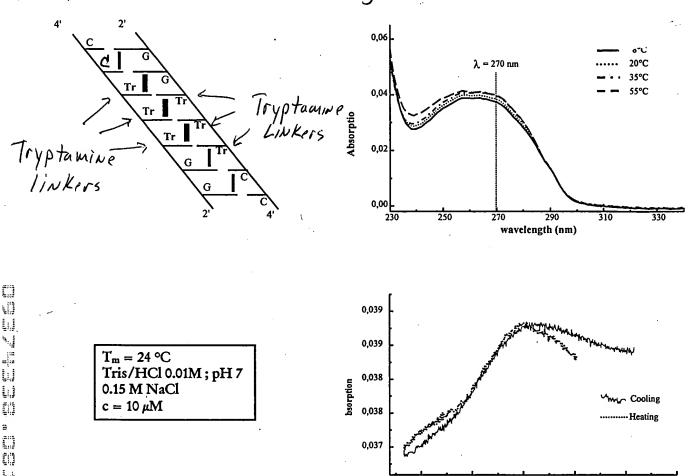






 $T_{m} = 61 \, ^{\circ}\mathbb{C}$ $T_{m} = 61 \, ^{\circ}\mathbb{C}$ $0.15 \, M_{m} \, M_{a} \, \text{Cl}$ $c = 5 \, \mu M + 5 \, \mu M + 5 \, \mu M$

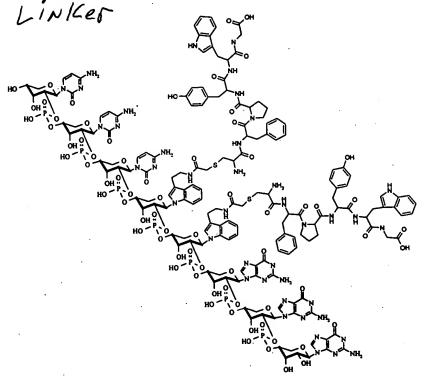
Figure 23- p-RNA Sequence with Multiple Tryptamine Linkers

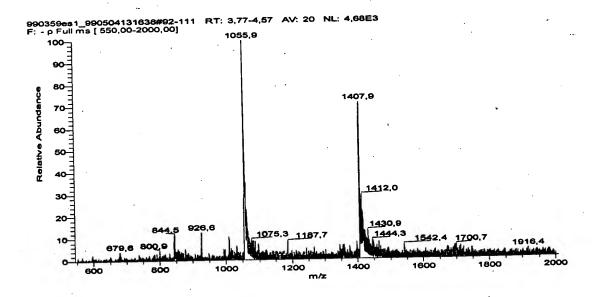


wavelength (nm)

Thermodynamic data of the self pairing p-RNA oligomer 4' CCTrTrTrGG 2'

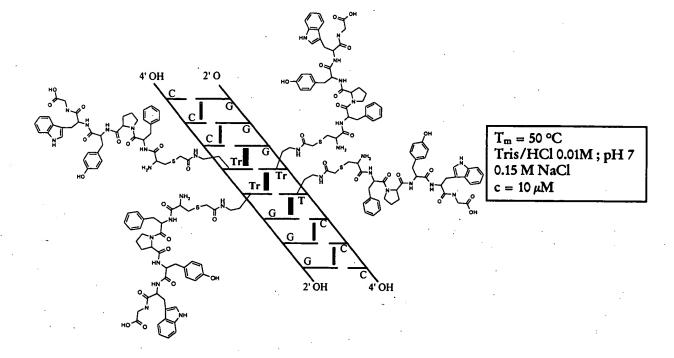
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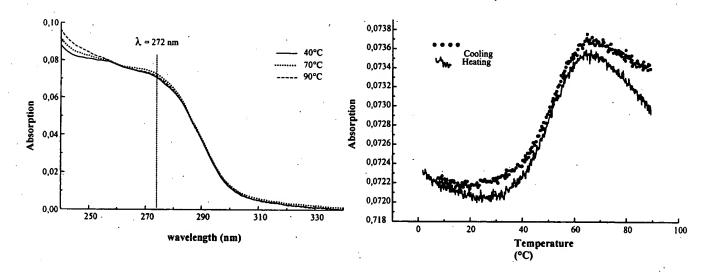




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Figure 25 P-RNA Hybrid Tructure With Four Peptides

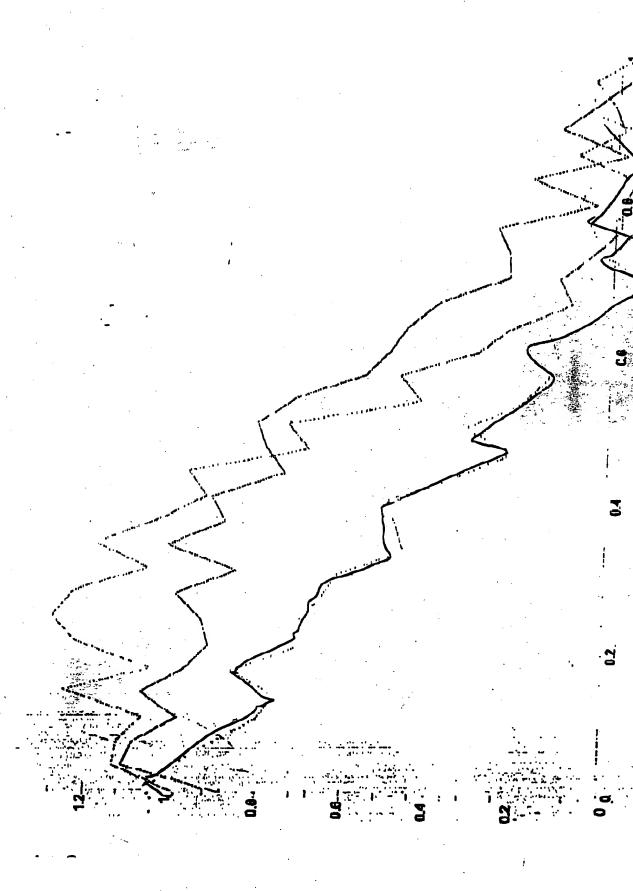


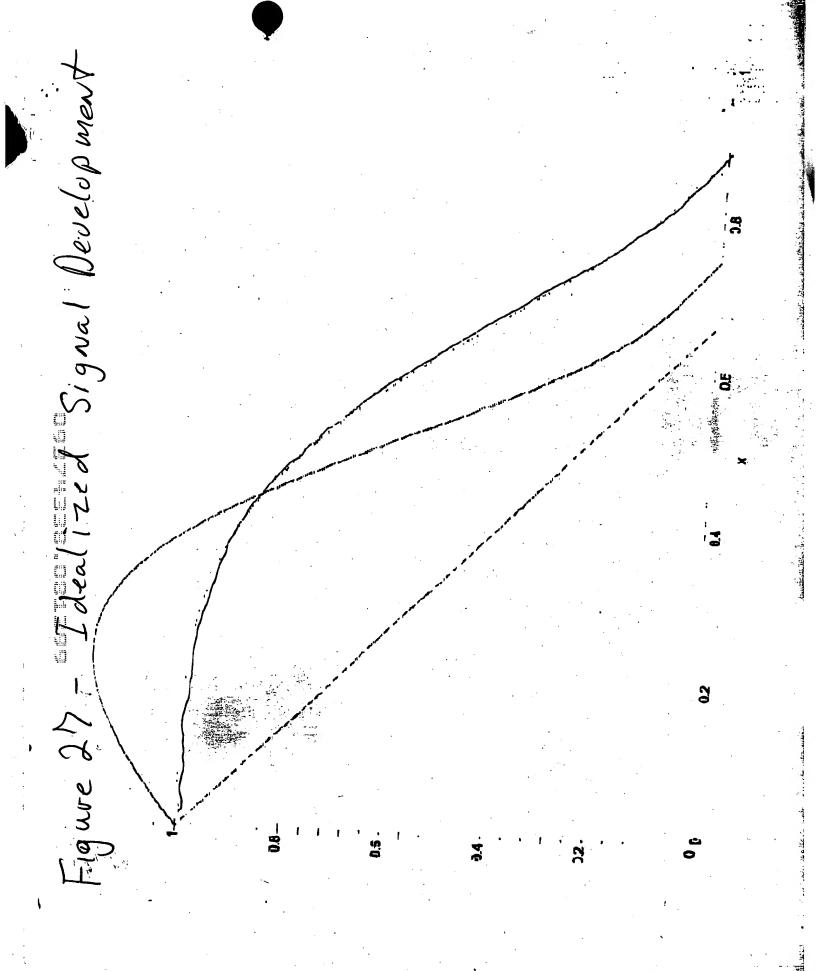


Thermodynamic data of the p-RNA peptide conjugate

£ 15

s-Simulation of Measurements With Noise





St und 1

10 000 different chip position events with a simulation of Factoranalysis clustering of the signal development on noise of absolute +/- 25 % (50%).

